Cqc

Attorney Docket Number: QA211

DEC 1 2 2005

CERTIFICATE OF MAILING

Dereby certify that this paper (along with any paper referred to as being attached or enclosed) is being deposited with the United States Postal Service on the date shown below with sufficient postage as first class mail in an envelope addressed to the: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

1450, Alexandria, VA 22313-1450.

Type or print name

Signature Signature

Dec 8,2005

### IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

IN RE APPLICATION OF

BANVILLE ET AL.

**APPLICATION NO: 09/848,694** 

(now U.S. Pat. 6,924,391)(32)

FILED: MAY 3, 2001

FOR: ALPHA-AMINO, -THIO, -OXO SUBSTITUTED KETONES AS

PHOSPHOLIPASE INHIBITORS

Certificate

DEC 1 5 2005

of Correction

Commissioner for Patents

P.O. Box 1450

Alexandria, VA 22313-1450

ATTN: Decision and Certification Branch of Patent Issue Division

### REQUEST FOR CERTIFICATE OF CORRECTION UNDER 37 C.F.R. 1.322

Sir:

The above patent application Serial No. 09/848,694 issued as U.S. Pat. No. 6,924,391 on August 2, 2005. Applicants hereby request a certificate of correction for the errors set forth on form PTO/SB/44 attached hereto as Appendix A. Remarks regarding the corrections are set forth below on pages 2-4.

### **REMARKS**

For the USPTO's convenience, Applicants have attached the following appendices:

- A completed form PTO/SB/44 listing all the corrections sought to be made herein (Appendix A).
- Copies of the Office Action mailed July 8, 2004, the Amendment filed October 7,
   2004 and the Notice of Allowance (Appendix B).

Each of the errors, depicted in Table 1, occurred through the fault of the USPTO, for which correction is sought under 37 C.F.R. 1.322. Applicants submit these corrections present no new matter and do not require further examination. Remarks regarding each of the corrections follow the Table.

Table 1

Error	Column	Error	Correction
Number and Page			
1	Col. 119,	Numerous compounds are missing	After "A compound selected from",
	line 2	from claim 1.	the following formulae should be
			added:
			CH <sub>3</sub>

Attorney	Dooleat	Mumba	OA211	
Auomev	Docker	ivumbe	r: UAZTI	

7 Ittorney De	ocket Number:	QAZII	
			CH <sub>3</sub> CH <sub>3</sub> CH <sub>3</sub> CH <sub>3</sub> CH <sub>3</sub> CO <sub>2</sub> H And CO <sub>2</sub> H And CO <sub>2</sub> H And CH <sub>3</sub> CH <sub>3</sub> CO <sub>2</sub> H And CO
2	Col. 119,	The structure is incorrect	Replace the printed structure with
	lines 45-65		the following:
3	Col. 120	The etmetions is in a second	CH <sub>3</sub> COCONH CCO <sub>2</sub> H
3	Col. 120,	The structure is incorrect	Replace the printed structure with
	lines 25-45		the following:
4	Col. 122,	The structure is incorrect	Replace the printed structure with
	lines 25-45		the following:
			CI CH3 CONHCH2CH2CH

US Pat. No. 6, 924,391 (application Serial No. 09/848,694)

Attorney Docket Number: QA211

7 titoliney D	ocket Number:	QA211	
5	Col. 122,	The structure is incorrect	Replace the printed structure with
	lines 45-65		the following:
		·	CI CO H
6	Col. 125,	The structure is incorrect	Replace the printed structure with
	lines 20-40		the following:
			CH <sub>3</sub> CO <sub>2</sub> H

In Error 1, nine compounds were omitted from claim 1. These compounds were allowed in claim 11 of Applicants' Amendment dated October 7, 2004. Applicants submit that the nine unmarked compounds shown on page 10 of the Amendment have been omitted from claim 1 of the patent and should be added to the claim following the text "A compound selected from."

Errors 2-6 consist of five structures incorrectly printed. All are USPTO mistakes. This is apparent in comparing the structures found on pages 11-12 of Applicants' Amendment with the printed patent.

As each of the mistakes made are the fault of the USPTO, Applicants believe no fees are required. If a fee is deemed to be required, the Commissioner is hereby authorized to charge such fee to Deposit Account No. 19-3880. The USPTO is requested to kindly contact the undersigned if deemed appropriate to expedite this request.

<sup>&</sup>lt;sup>1</sup> On page 3 of the Office Action, mailed July 8, 2004, the Examiner states that "Claim 11 will be allowed to the extent it reads on the elected subject matter. Compounds containing Silicon and heterocyclic subject matter should be deleted." As shown on pages 9-13 of Applicants' Amendment, claim 11 was amended to remove silicon and heterocyclic subject matter which reflected the scope of the generic concept of the elected subject matter. On page 2 of the Notice of Allowance, the Examiner states that "specific compounds of claim 11 are now claimed, along with the pharmaceutical composition. These compounds do not read on the above reference and hence are patentable."

US Pat. No. 6, 924,391 (application Serial No. 09/848,694) Attorney Docket Number: QA211

Respectfully submitted,

Pamela A. Mingo, Ph.D.

Agent for Applicant

Reg. No. 48,256

Bristol-Myers Squibb Company Patent Department P.O. Box 4000 Princeton, NJ 08543-4000 (203) 677-7669

Date: December 8, 2005

- 5 -



# APPENDIX A

### **CERTIFICATE OF CORRECTION**

PATENT NO : 6,924,391 \( \beta \) 2
DATED: : August 2, 2005

INVENTOR(S) : Jacques Banville, Roger Remillard, Neelakantan Balasubramanian, Gilles

Bouthillier, Alain Martel

It is certified that an error appears or errors appear in the above-identified patent and that said Letters Patent is hereby corrected as shown below:

Col. 119, line 2, after "A compound selected from", the following formulae should be added:

MAILING ADDRESS OF SENDER:

PATENT NO. 6,924,391

Pamela A. Mingo, Ph.D.
Bristol-Myers Squibb Company
Patent Department
P.O. Box 4000
Princeton, NJ 08543-4000
(203) 677-7669

Page 1 of 4

FORM PTO/SB/44

### **CERTIFICATE OF CORRECTION**

PATENT NO

6,924,391 B2

DATED:

August 2, 2005

INVENTOR(S)

Jacques Banville, Roger Remillard, Neelakantan Balasubramanian, Gilles

Bouthillier, Alain Martel

### (Continued)

## Col. 119, lines 45-65, the formula should appear as follows:

MAILING ADDRESS OF SENDER:

PATENT NO. 6,924,391

Pamela A. Mingo, Ph.D. Bristol-Myers Squibb Company Patent Department P.O. Box 4000 Princeton, NJ 08543-4000 (203) 677-7669

Page 2 of 4

# **CERTIFICATE OF CORRECTION**

PATENT NO

6,924,391 62

DATED:

August 2, 2005

INVENTOR(S)

Jacques Banville, Roger Remillard, Neelakantan Balasubramanian, Gilles

Bouthillier, Alain Martel

(Continued)

Col. 120, lines 25-45, the formula should appear as follows:

Col. 122, lines 25-45, the formula should appear as follows:

MAILING ADDRESS OF SENDER:

PATENT NO. 6,924,391

Pamela A. Mingo, Ph.D. Bristol-Myers Squibb Company Patent Department P.O. Box 4000 Princeton, NJ 08543-4000 (203) 677-7669

Page 3 of 4

FORM PTO/SB/44

### **CERTIFICATE OF CORRECTION**

PATENT NO

6,924,391 B 2

DATED:

August 2, 2005

INVENTOR(S)

Jacques Banville, Roger Remillard, Neelakantan Balasubramanian, Gilles

Bouthillier, Alain Martel

### (Continued)

Col. 122, lines 45-65, the formula should appear as follows:

Col. 125, lines 20-40, the formula should appear as follows:

MAILING ADDRESS OF SENDER:

PATENT NO. 6,924,391

Pamela A. Mingo, Ph.D.
Bristol-Myers Squibb Company
Patent Department
P.O. Box 4000
Princeton, NJ 08543-4000
(203) 677-7669

Page 4 of 4



# APPENDIX B

United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS COMMERCE

P.O. Box 1450 Alexandria, Virginia 22313-1450 www.uspto.gov

# D FEE(S) DUE BMS PATENT LAW 10/22/2004

Marla J. Mathias Bristol-Myers Squibb Company

Patent Department P.O. Box 4000

Princeton, NJ 08543-4000

OCT 26 2004

**EXAMINER** 

KUMAR, SHAILENDRA

Docketed Item

Due Date

ART UNIT 1621

AILED: 10/22/2004

PAPER NUMBER

Attorney [ APPLICATION NO. FILING DATE ^ 09/848,694· ~

05/03/2001

FIRST NAMED INVENTOR

ATTORNEY DOCKET NO. QA211

CONFIRMATION NO. 7932

TITLE OF INVENTION: ALPHA-AMINO,-THIO,-OXO SUBSTITUTED KETONES AS PHOSPHOLIPASE INHIBITORS

APPLN. TYPE nonprovisional	SMALL ENTITY NO	ISSUE FEE	PUBLICATION FEE	TOTAL FEE(S) DUE	
THE APPLICATION I	DENTIFIED ABO	VE HAS BEEN EVAN	\$300	\$1670	01/24/2005

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT. PROSECUTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN THREE MONTHS FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. STATUTORY PERIOD CANNOT BE EXTENDED. SEE 35 U.S.C. 151. THE ISSUE FEE DUE INDICATED ABOVE REFLECTS A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE APPLIED IN THIS APPLICATION AN EQUIVALENT) MUST BE RETURNED WITHIN THIS PERIOD EVEN IF NO FEE IS DE 85B (OR HOW TO REPLY TO THIS NOTICE:

I. Review the SMALL ENTITY status shown above.

If the SMALL ENTITY is shown as YES, verify your current SMALL ENTITY status:

A. If the status is the same, pay the TOTAL FEE(S) DUE shown

B. If the status above is to be removed, check box 5b on Part B -Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and twice the amount of the ISSUE FEE shown above, or

NOV 03 2004

If the SMALL ENTITY is show RATEN DEPARTMENT WALLINGFORD, CT BRISTOL-MYERS SQUIBB COMPANY

A. Pay TOTAL FEE(S) DUE shown above, or

B. If applicant claimed SMALL ENTITY status before, or is now claiming SMALL ENTITY status, check box 5a on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and 1/2

II. PART B - FEE(S) TRANSMITTAL should be completed and returned to the United States Patent and Trademark Office (USPTO) with your ISSUE FEE and PUBLICATION FEE (if required). Even if the fee(s) have already been paid, Part B - Fee(s) Transmittal should be completed and returned. If you are charging the fee(s) to your deposit account, section "4b" of Part B - Fee(s) Transmittal should be

[II. All communications regarding this application must give the application number. Please direct all communications prior to issuance to

MPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of naintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.





### United States Patent and Trademark Office

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Box 1450 Alexandria, Virginia 22313-1450 www.uspto.gov

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/848,694	05/03/2001	P E Jacques Banville	QA211	7932
759	10/22/2004	As a	EXAM	INER
Marla J. Mathias Bristol-Myers Squil	bb Company	DEC 1 2 2005 w	KUMAR, SH	AILENDRA
Patent Department	·	\o \&/	ART UNIT	PAPER NUMBER
P.O. Box 4000 Princeton, NJ 08543	-4000	TRADEMARKS.	1621	
, , , , , , , , , , , , , , , , , , , ,		HADEM	DATE MAILED: 10/22/2004	1

### Notice of Fee Increase on October 1, 2004

If a reply to a "Notice of Allowance and Fee(s) Due" is filed in the Office on or after October 1, 2004, then the amount due will be higher than that set forth in the "Notice of Allowance and Fee(s) Due" because some fees will increase effective October 1, 2004. See Revision of Patent Fees for Fiscal Year 2005; Final Rule, 69 Fed. Reg. 52604, 52606 (May 10, 2004).

The current fee schedule is accessible from WEB site (http://www.uspto.gov/main/howtofees.htm).

If the fee paid is the amount shown on the "Notice of Allowance and Fee(s) Due" but not the correct amount in view of the fee increase, a "Notice of Pay Balance of Issue Fee" will be mailed to applicant. In order to avoid processing delays associated with mailing of a "Notice of Pay Balance of Issue Fee," if the response to the Notice of Allowance is to be filed on or after October 1, 2004 (or mailed with a certificate of mailing on or after October 1, 2004), the issue fee paid should be the fee that is required at the time the fee is paid. See Manual of Patent Examining Procedure (MPEP), Section 1306 (Eighth Edition, Rev. 2, May 2004). If the issue fee was previously paid, and the response to the "Notice of Allowance and Fee(s) Due" includes a request to apply a previously-paid issue fee to the issue fee now due, then the difference between the issue fee amount at the time the response is filed and the previously-paid issue fee should be paid. See MPEP Section 1308.01.

Effective October 1, 2004, 37 CFR 1.18 is amended by revising paragraphs (a) through (c) to read as set forth below.

Section 1.18 Patent post allowance (including issue) fees.

(a) Issue fee for issuing each original or reissue patent, except a design or plant patent:

(b) Issue fee for issuing a design patent:

(c) Issue fee for issuing a plant patent:

By a small entity (Sec. 1.27(a))......\$330.00

By other than a small entity......\$660.00

Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at (703) 305-8283.

OIPE	Application No.	Applicant(s)
Notice of Allowability	09/848,694	
	Examiner	BANVILLE ET AL.
DEC 1 2 2005	//	Art Unit
	SHAILENDRA - KUMAR	1621
The MAILING DATE of this compared appears appeared to previously mailed), a Notice of Allowance (PTOL-85) NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RIOT the Office or upon petition by the applicant. See 37 CFR 1.313  1. This communication is responsive to 10/7/04.	ears on the cover sheet with the (OR REMAINS) CLOSED in this or other appropriate communical IGHTS. This application is subject and MPEP 1308.	e correspondence address application. If not included tion will be mailed in due course. The ct to withdrawal from issue at the ini
2. The allowed claim(s) is/are 11 and 9(renumbered as 1-2).		
3. The drawings filed on are accepted by the Examiner.		
4. LJ Acknowledgment is made of a claim for foreign priority and	4	
4. ☐ Acknowledgment is made of a claim for foreign priority und  a) ☐ All b) ☐ Some* c) ☐ None of the:	aer 35 U.S.C. § 119(a)-(d) or (f).	
1. Certified copies of the priority documents.		
3. Copies of the certified copies of the artistic	peen received in Application No.	
International Bureau /PCT Duta 47 Page 17	ments have been received in this	s national stage application
* Certified copies not received:		application from the
Applicant has THREE MONTHS FROM THE "MAILING DATE" of oted below. Failure to timely comply will result in ABANDONMEN HIS THREE-MONTH PERIOD IS NOT EXTENDABLE.  A SUBSTITUTE OATH OR DECLARATION must be submitted INFORMAL PATENT APPLICATION (PTO-152) which gives replacement above.	d. Note the attached EXAMINER	•
· · · · - O · LD DRAWINGS / 66 #		The actional fit
	s Patant Dawin -	
(a) ☐ including changes required by the Notice of Draftsperson's  1) ☐ hereto or 2) ☐ to Paper No./Mail Date	s ratent Drawing Review (PTO-9	948) attached
(b) ☐ including changes required by the attached Examiner's An Paper No./Mail Date  dentifying indicia custs as the	nand	
Paper No./Mail Date	nenament / Comment or in the Ot	ffice action of
each sheet. Replacement sheet(s) should be labeled as such in the he	)) should be written on the drawing	gs in the front (not the back) of
attached Examiner's comment regarding REQUIREMENT FOR	THE DEPOSIT OF BIOLOGICAL	ust be submitted. Note the L MATERIAL.
hment(s)	•	
Notice of References Cited (PTO-892)	5 D Notice at a	
Notice of Draftperson's Patent Drawing Review (PTO-948)	5. Notice of Informal Pate	ent Application (PTO-152)
nformation Disclosure Statements (PTO-1449 or PTO/SR/09)	<ol> <li>Interview Summary (P' Paper No./Mail Date _</li> <li>⊠ Examiner's Amendmer</li> </ol>	TO-413),
xaminer's Comment Regarding Requirement for Deposit Biological Material	8. ⊠ Examiner's Statement of	
ent and Trademark Office	y. [_] Other	·

Art Unit: 1621

1. An examiner's amendment to the record appears below. Should the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the issue fee.

Authorization for this examiner's amendment was given in a telephone interview with Pamela Mingo on 10/19/04.

The application has been amended as follows:

- 1) Claims 1-4, 7, 10, 12 and 13 have been canceled without prejudice.
- 2) In claim 9, line 3, "1" is changed to - 11 -.
- 2. The following is an examiner's statement of reasons for allowance: The closest prior art was JP 5-222006, especially page 5, top compound. However with the cancellation of above claims, specific compounds of claim 11 are now claimed, along with the pharmaceutical composition. These compounds do not read on the above reference and hence are patentable.

Any comments considered necessary by applicant must be submitted no later than the payment of the issue fee and, to avoid processing delays, should preferably accompany the issue fee. Such submissions should be clearly labeled "Comments on Statement of Reasons for Allowance."

3. Any inquiry concerning this communication or earlier communications from the examiner should be directed to SHAILENDRA - KUMAR whose telephone number is (571)272-0640. The examiner can normally be reached on Mon-Thur 8:00-5:30, Alt Fri.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Johann Richter can be reached on (571)272-0646. The fax phone number for the organization where this application or proceeding is assigned is 703-872-9306.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).

SHAILENDRA - KUMAR Primary Examiner Art Unit 1621

S.Kumar 10/19/04



# United States Patent and Trademark Office

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Box 1450 Alexandria, Virginia 223 13-1450

			www.uspto.gov	
APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR		
09/848,694	05/03/2001		ATTORNEY DOCKET NO.	CONFIRMATION NO.
	03/03/2001	Jacques Banville	QA211	7932
759	0 10/22/2004			
Marla J. Mathias			EXAMI	NER
Bristol-Myers Squil	ob Company		KUMAR, SHA	AILENDRA
Patent Department				
P.O. Box 4000			ART UNIT	PAPER NUMBER
Princeton, NJ 08543	-4000		1621	
			DATE MAILED: 10/22/2004	

# Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 367 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 367 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (703) 305-1383. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at (703) 305-8283.

PLK " Monts



### United States Patent and Trademark Office

QADZII US-NP

UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Box 1450 Alexandria, Virginia 22313-1450

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
09/848,694	05/03/2001	Jacques Banville	QA211	7932
75	90 07/08/2004		EXAM	INER
Marla J. Mathi	as		KUMAR, SH	AILENDRA
Bristol-Myers S	Squibb Company			
Patent Departme			ART UNIT	PAPER NUMBER
P.O. Box 4000			1621	· · · · · · · · · · · · · · · · · · ·

AUG 12 2004

DATE MAILED: 07/08/2004

BRISTOL MUSE SIGNIFIED AND PARTIES OF SERVICE ALERA WALLINGFORD, OT

SENI (SC)

Please find below and/or attached an Office communication concerning this application or proceeding.



# RECEIVED BNS PATENT LAW

JUL 12 2004

Docketed Item First CA

Due Date

Attorney VOLLES

# FINAL REJECTION

RECEIVED

Princeton, NJ 08543-4000

RECEIVED

JUL 2 9 2004

JUL 2 9 2014

Woodcock Washburn

DOCKET DEPT

Final Review 9/8/04 Final Reject 10/8/04 Notice of App 10/8/04

TECH VIII

AUG 12, 2004

BRISTOL-MYERS SQUUE COMPANY
PATENT DEPARTMENT
WALLINGFORD, CT



	Application No.	Applicant(s)
Office Action Summary	09/848,694	BANVILLE ET AL.
Janmary	Examiner	Art Unit
The MAILING DATE of this	SHAILENDRA - KUMAR	
The MAILING DATE of this communication Period for Reply	appears on the cover sheet with	the correspondence address
A SHORTENED STATUTORY PERIOD FOR REF THE MAILING DATE OF THIS COMMUNICATION  Extensions of time may be available under the provisions of 37 CFR after SIX (6) MONTHS from the mailing date of this communication.  If the period for reply specified above is less than thirty (30) days, a r  If NO period for reply is specified above, the maximum statutory perior  Failure to reply within the set or extended period for reply will, by stat Any reply received by the Office later than three months after the mai earned patent term adjustment. See 37 CFR 1.704(b).  Status  1) Responsive to communication(s) filed on 29  2a) This action is FINAL.  2b) Th  3) Since this application is in condition for allow closed in accordance with the practice under closed in accordance with the practice under Disposition of Claims  4) Claim(s) 1-4,7 and 9-13 is/are pending in the 4a) Of the above claim(s) 10 is/are withdrawn  5) Claim(s) 11 is/are allowed.  Claim(s) 1-4,9,12 and 13 is/are rejected.	1.136(a). In no event, however, may a reply eply within the statutory minimum of thirty (30 and will apply and will expire SIX (6) MONTHS ute, cause the application to become ABAND liling date of this communication, even if timely application is non-final.  April 2004.  This action is non-final.  Ance except for formal matters, Ex parte Quayle, 1935 C.D. 11, applicable.	be timely filed  i) days will be considered timely.  from the mailing date of this communication.  ONED (35 U.S.C. § 133).  y filed, may reduce any
8) Claim(s) Z is/are objected to. 8) Claim(s) are subject to restriction and/o	or election requirement.	
9) The specification is objected to by the Examine 10) The drawing(s) filed on is/are: a) access applicant may not request that any objection to the of Replacement drawing sheet(s) including the correction of the oath or declaration is objected to by the Examine that the control of the oath or declaration is objected to by the Examine that the control of the oath or declaration is objected to by the Examine that the control of the oath or declaration is objected to by the Examine that the control of the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to by the Examine that the oath or declaration is objected to be obj	epted or b) objected to by the drawing(s) be held in abeyance. So on is required if the drawing(s) is olearniner. Note the attached Office	ee 37 CFR 1.85(a). bjected to. See 37 CFR 1.121(d). e Action or form PTO-152.
1. Certified copies of the priority documents 2. Certified copies of the priority documents 3. Copies of the certified copies of the priority application from the International Bureau ( * See the attached detailed Office action for a list of the certified copies of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the certified copies of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the certified copies of the priority documents application from the International Bureau (  * See the attached detailed Office action for a list of the certified copies of the priority documents are copied to the priority documents are copied to the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the International Bureau (  * See the attached detailed Office action for a list of the priority application from the Internation for a list of the priority application from the Internation for a list of the Internation for a	have been received. have been received in Application y documents have been received (PCT Rule 17.2(a)). The certified copies not received  4)  Interview Summary (6)	on No ed in this National Stage d.
Information Dicalogue 91	Paper No(s)/Mail Date	- '''

Application/Control Number: 09/848,694

Art Unit: 1621

## **DETAILED ACTION**

This office action is in response to applicants' communication filed on 4/29/04. Claims 1-4, 7, and 9-13 are pending in this application. Claims 5-6 and 8 have been canceled. Claim 10 has been withdrawn from the consideration, being drawn to the non-elected invention.

Rejection of claims 1-4 and 9 over Kun et al is hereby withdrawn subsequent to applicants' amendment.

# Claim Rejections - 35 USC § 102

1. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless -

- (b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.
- 2. Claims 1-4, 9, 12 and 13 are rejected under 35 U.S.C. 102(b) as being anticipated by JP 5-222006.

JP'006, page 5, lines 1-5, anticipate instant claims when, A is halo, R3, R4, R5 are H, R2 is H, W is O, and R1 is alkoxy, see page 9, Table. Inasmuch, the compound is soluble in water, the composition is anticipated. English abstract is attached along with the Japanese document.

3. Claims 7 is objected to as being dependent upon a rejected base claim, but would be allowable to the extent that it reads on the elected subject matter, if rewritten in independent form including all of the limitations of the base claim and any

Application/Control Number: 09/848,694

Art Unit: 1621

intervening claims. Note applicants should delete Silicon containing subject matter.

- Claim 11 will be allowed to the extent it reads on the elected subject matter.
   Compounds containing Silicon and heterocyclic subject matter should be deleted.
- 5. Applicant's amendment necessitated the new ground(s) of rejection presented in this Office action. Accordingly, **THIS ACTION IS MADE FINAL**. See MPEP § 706.07(a). Applicant is reminded of the extension of time policy as set forth in 37 CFR 1.136(a).

A shortened statutory period for reply to this final action is set to expire THREE MONTHS from the mailing date of this action. In the event a first reply is filed within TWO MONTHS of the mailing date of this final action and the advisory action is not mailed until after the end of the THREE-MONTH shortened statutory period, then the shortened statutory period will expire on the date the advisory action is mailed, and any extension fee pursuant to 37 CFR 1.136(a) will be calculated from the mailing date of the advisory action. In no event, however, will the statutory period for reply expire later than SIX MONTHS from the date of this final action.

6. Any inquiry concerning this communication or earlier communications from the examiner should be directed to SHAILENDRA - KUMAR whose telephone number is (571)272-0640. The examiner can normally be reached on Mon-Thur 8:00-5:30, Alt Fri.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Johann Richter can be reached on (571)272-0646. The fax phone number for the organization where this application or proceeding is assigned is 703-872-9306.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).

SHAILENDRA - KUMAR Primary Examiner Art Unit 1621

S.Kumar 7/6/04 Notice of Reference Cites

Application/Control No. Applicant(s)/Patent Under 09/848,694 Reexamination BANVILLE ET AL. Examiner Art Unit SHAILENDRA - KUMAR Page 1 of 1 1621

*	l	Document Number	Date	U.S. PATENT DOCUMENTS	
		Country Code-Number-Kind Code	MM-YYYY	Name	 
	A.	US-		ivame :	 Classification
	В	US-			
	С	US-			
$\perp$	D	US-			
	E	US-		*	
	F	US-			
$\perp$	G	US-			
$\int$	н	US-	<u> </u>		
	1	US-			
	J	US-			1
	ĸ	US-			
L	L	US-			
_ N	N	US-			
			For		
		Document Number Country Code-Number Kind Code	Date	EIGN PATENT DOCUMENTS	

*		Document Number	Date	FOREIGN PATENT DO	DCUMENTS	
	<del> </del>	Country Code-Number-Kind Code	MM-YYYY	Country		<del></del>
<u> </u>	N	5-222006	08-1993	Japan	Name	Classification
	0				<u> </u> -	
	P					
	Q					-
	R					
	s					
	Т					
*				NON-PATENT DOCUM		

*	NON-PATENT DOCUMENTS	
+	Include as applicable: Author, Title Date, Publisher, Edition or Volume, Pertinent Pages)	
U		
V		
w		
x		
Py of this in MM-	reference is not being furnished with this Office action. (See MPEP § 707.05(a).)	

Copy of this reference is not being furnished with this Office action. (See MPEP § 707.05(a).) ates in MM-YYYY format are publication dates. Classifications may be US or foreign.



#### CERTIFICATE OF MAILING

I hereby certify that this paper (along with any paper referred to as being attached or enclosed) is being deposited with the United States Postal Service on the date shown below with sufficient postage as first class mail in an envelope addressed to the: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

Pamela A. Mingo

Type or print name

October 7, 2004

Date

# IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of: Banville et. al

Art Unit: 1621

Application No.: 09/848,694

Examiner: S. Kumar

Filed: May 3, 2001

For: Alpha-Amino, -Thio, -Oxo Substituted Ketones as Phopholipase Inhibitors

MAIL STOP: Amendment Commissioner for Patents

P.O. Box 1450

Alexandria, VA 22313-1450

### REPLY AND AMENDMENT AFTER FINAL PURSUANT TO 37 C.F.R. §1.116 Sir:

This Reply and Amendment is a timely response to a Final Rejection dated. July 8, 2004 having a three-month shortened statutory period for response expiring October 8, 2004. In Examiner's first Office Action dated January 5, 2004, Claims 1-4 and 9 were rejected and Claims 7 and 8 were objected to as being dependent upon a rejected base claim. Applicants canceled Claims 5,6, and 8, amended Claims 1-3, 7, and 9, and added Claims 11, 12, and 13. Applicants respectfully request reconsideration of the above-entitled application in light of the following amendments and remarks. Applicants note that a NOTICE OF APPEAL has been filed in concert with the present Reply and Amendment.

Amendments to the Claims are reflected in the listing of claims which begins on page 2 of this paper.

Remarks/Arguments begin on page 18 of this paper.

### Amendments to the Claims:

This listing of claims will replace all prior versions and listings of claims in the application:

### **Listing of Claims:**

Claim 1. (Currently Amended) A compound of the formula

wherein  $X_1$  is O,  $S(O)_n$ ,  $-\stackrel{R^5}{N}$ ,  $co-\stackrel{R^5}{N}$ , or -CH<sub>2</sub>-, with the proviso that when  $X_1$  is -CH<sub>2</sub>-,  $R^1$  and  $R^2$  are only halogen;

n is 0, 1 or 2;

 $R^a$  and  $R^b$  when taken together form an oxo (=O) group, or  $R^a$  and  $R^b$  are each independently hydrogen, OH, OCOR<sup>9</sup>, NH<sub>2</sub>, N<sub>3</sub>, NHCOOR<sup>9</sup>, NHCOCOR<sup>9</sup>, NHSO<sub>2</sub>R<sup>9</sup> or F;

X is H, CF<sub>3</sub>, OCF<sub>3</sub>, halogen, C<sub>1</sub>–C<sub>7</sub> alkyl, C<sub>2</sub>–C<sub>7</sub> alkenyl, C<sub>2</sub>–C<sub>7</sub> alkynyl or C<sub>3</sub>–C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by COOR<sup>8</sup>, CN, C(O)NR<sup>6</sup>R<sup>7</sup>, PO<sub>3</sub>R<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, heterocyclic, OR<sup>8</sup>, SH, S(O)<sub>n</sub>R<sup>9</sup>, NR<sup>6</sup>R<sup>7</sup>, NH(CO)NR<sup>6</sup>R<sup>7</sup>, NH(CO)OR<sup>9</sup>, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from NR<sup>6</sup>R<sup>7</sup>, OR<sup>8</sup>, COOR<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, OCOR<sup>9</sup>, PO<sub>3</sub>R<sup>8</sup>, and C(O)NR<sup>6</sup>R<sup>7</sup> and heterocyclic;

R<sup>1</sup> and R<sup>2</sup> are each independently H, halogen, OR<sup>9</sup>, C<sub>1</sub>-C<sub>7</sub> alkyl, C<sub>2</sub>-C<sub>7</sub> alkynyl,

C2–C7 alkenyl or C3–C7 cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by COOR<sup>8</sup>, CN, C(O)NR<sup>6</sup>R<sup>7</sup>, PO<sub>3</sub>R<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, heterocyclie, OR<sup>8</sup>, SH, S(O)<sub>n</sub>R<sup>9</sup>, NR<sup>6</sup>R<sup>7</sup>, NH(CO)NR<sup>6</sup>R<sup>7</sup>, NH(CO)OR<sup>9</sup>, OC(O)OR<sup>9</sup>, or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted with one or two groups independently selected from NR<sup>6</sup>R<sup>7</sup>, OR<sup>8</sup>, COOR<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, OCOR<sup>9</sup>, PO<sub>3</sub>R<sup>8</sup>, and C(O)NR<sup>6</sup>R<sup>7</sup> and heterocyclie;

 $R^3$ ,  $R^4$  and Y are each independently H, halogen,  $OR^{10}$ ,  $S(O)_nR^{10}$ ,  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by  $COOR^8$ , CN,  $C(O)NR^6R^7$ ,  $PO_3R^8$ ,  $SO_3R^8$ , heterocyclic,  $OR^8$ , SH,  $S(O)_nR^9$ ,  $NR^6R^7$ ,  $NH(CO)NR^6R^7$ ,  $NH(CO)OR^9$ ,  $OC(O)OR^9$ , or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted by one or two groups independently selected from  $NR^6R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3R^8$ ,  $OCOR^8$ ,  $PO_3R^8$ , and  $C(O)NR^6R^7$  and heterocyclic, with the proviso that not all of  $R^3$ ,  $R^4$  and Y may be the same halogen;

R<sup>5</sup>, R<sup>6</sup> and R<sup>7</sup> are each independently H, C<sub>1</sub>–C<sub>7</sub> alkyl, C<sub>2</sub>–C<sub>7</sub> alkenyl, C<sub>2</sub>–C<sub>7</sub> alkynyl or C<sub>3</sub>–C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by COOR<sup>8</sup>, CN, OR<sup>8</sup>, NR<sup>8</sup>R<sup>9</sup>, SO<sub>3</sub>R<sup>8</sup>, PO<sub>3</sub>R<sup>8</sup>, halogen, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from COOR<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, and PO<sub>3</sub>R<sup>8</sup> and heterocyclie;

R<sup>8</sup> is H, C<sub>1</sub>-C<sub>7</sub> saturated straight chain alkyl or cycloalkyl;

R<sup>9</sup> is C<sub>1</sub>-C<sub>7</sub> saturated straight chain alkyl or cycloalkyl;

 $R^{10}$  is  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl, aryl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, aryl or cycloalkyl group being optionally substituted by

COOR<sup>8</sup>, CN, C(O)NR<sup>6</sup>R<sup>7</sup>, PO<sub>3</sub>R<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, heterocyclic, OR<sup>8</sup>, SH, S(O)<sub>n</sub>R<sup>9</sup>, NR<sup>6</sup>R<sup>7</sup>, NH(CO)NR<sup>6</sup>R<sup>7</sup>, NH(CO)OR<sup>9</sup>, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from NR<sup>6</sup>R<sup>7</sup>, OR<sup>8</sup>, COOR<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, OCOR<sup>8</sup>, PO<sub>3</sub>R<sup>8</sup>, and C(O)NR<sup>6</sup>R<sup>7</sup> and heterocyclic;

Z is  $OR^{11}$ ,  $S(O)_nR^{11}$ ,  $NR^{11}R^{12}$  or  $CHR^{11}R^{12}$ ;

 $R^{11}$  is  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being substituted by  $NR^{13}R^{14}$ ,  $S(O)_nR^{13}$ , or  $OR^{13}$ ;

 $R^{12}$  is hydrogen,  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by  $NR^{13}R^{14}$ ,  $S(O)_nR^{13}$ , or  $OR^{13}$ ;

R<sup>13</sup> is SiR<sup>15</sup>R<sup>16</sup>R<sup>17</sup>, C<sub>1</sub>-C<sub>7</sub> alkyl, C<sub>2</sub>-C<sub>7</sub> alkenyl, C<sub>2</sub>-C<sub>7</sub> alkynyl, aryl or C<sub>3</sub>-C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being substituted by one to three groups independently selected from COOR<sup>8</sup>, OR<sup>8</sup>, Si R<sup>15</sup>R<sup>16</sup>R<sup>17</sup>, OR<sup>15</sup>, aryl, and biaryl and heteroaryl, said aryl[[,]] and biaryl and heteroaryl being optionally substituted with one to three groups independently selected from halogen, CF<sub>3</sub>, OR<sup>8</sup>, COOR<sup>8</sup>, NO<sub>2</sub>, and CN;

R<sup>14</sup> is H, SiR<sup>15</sup>R<sup>16</sup>R<sup>17</sup>, C<sub>1</sub>-C<sub>7</sub> alkyl, C<sub>2</sub>-C<sub>7</sub> alkenyl, C<sub>2</sub>-C<sub>7</sub> alkynyl, aryl or C<sub>3</sub>-C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by one to three groups independently selected from COOR<sup>8</sup>, OR<sup>8</sup>, Si R<sup>15</sup>R<sup>16</sup>R<sup>17</sup>, OR<sup>15</sup>, aryl, and biaryl and heteroaryl, said aryl[[,]] and biaryl and heteroaryl being optionally substituted with one to three groups independently selected from halogen, CF<sub>3</sub>, OR<sup>8</sup>, COOR<sup>8</sup>, NO<sub>2</sub>, and CN; and of

R<sup>13</sup> and R<sup>14</sup> when taken together with the nitrogen atom to which they are attached may form a 5—7 membered heterocyclic ring with one or more heteroatoms selected from O, N and S; said ring being optionally substituted by OR<sup>8</sup>, COOR<sup>8</sup>, or C(O)NR<sup>5</sup>R<sup>6</sup>; and

 $R^{15}$ ,  $R^{16}$ ,  $R^{17}$  are each independently is  $C_1$ - $C_7$  alkyl, aryl, benzyl, benzhydryl, biaryl, heteroaryl, or  $(C_1$ - $C_6)$  alkyl-aryl or  $(C_1$ - $C_6)$  alkyl-heteroaryl, said aryl, benzyl, benzhydryl, and biaryl being optionally substituted by halogen,  $CF_3$ ,  $OR^8$ ,  $COOR^8$ ,  $NO_2$ , CN, or  $C_1$ - $C_7$  alkyl.

### Claim 2. (Currently Amended) A compound of the formula

or a pharmaceutically acceptable salt thereof wherein

 $X_1$  is O,  $S(O)_n$ ,  $-\stackrel{R^5}{N}$ ,  $CO-\stackrel{R^5}{N}$  or -CH<sub>2</sub>-, with the proviso that when  $X_1$  is -CH<sub>2</sub>-,  $R^1$  and  $R^2$  are only halogen;

n is 0, 1 or 2;

 $R^a$  and  $R^b$  when taken together form an oxo (=0) group, or  $R^a$  and  $R^b$  are each independently hydrogen, OH, OCOR<sup>9</sup>, NH<sub>2</sub>, N<sub>3</sub>, NHCOOR<sup>9</sup>, NHCOCOR<sup>9</sup>, NHSO<sub>2</sub>R<sup>9</sup> or F;

X is H, CF<sub>3</sub>, OCF<sub>3</sub>, halogen, C<sub>1</sub>–C<sub>7</sub> alkyl, C<sub>2</sub>–C<sub>7</sub> alkenyl, C<sub>2</sub>–C<sub>7</sub> alkynyl or C<sub>3</sub>–C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by COOR<sup>8</sup>, CN, C(O)NR<sup>6</sup>R<sup>7</sup>, PO<sub>3</sub>R<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, heterocyclic, OR<sup>8</sup>, SH,

 $S(O)_n R^9$ ,  $NR^6 R^7$ ,  $NH(CO)NR^6 R^7$ ,  $NH(CO)OR^9$ , or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from  $NR^6 R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3 R^8$ ,  $OCOR^9$ ,  $PO_3 R^8$ , and  $C(O)NR^6 R^7$  and heterocyclic;

 $R^1$  and  $R^2$  are each independently H, halogen,  $OR^9$ ,  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkynyl,  $C_2$ – $C_7$  alkenyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by  $COOR^8$ , CN,  $C(O)NR^6R^7$ ,  $PO_3R^8$ ,  $SO_3R^8$ , heterocyclic,  $OR^8$ , SH,  $S(O)_nR^9$ ,  $NR^6R^7$ ,  $NH(CO)NR^6R^7$ ,  $NH(CO)OR^9$ ,  $OC(O)OR^9$ , aryl or heteroaryl, said aryl and heteroaryl being optionally substituted with one or two groups independently selected from  $NR^6R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3R^8$ ,  $OCOR^9$ ,  $PO_3R^8$ , and  $C(O)NR^6R^7$  and heterocyclic;

 $R^3$ ,  $R^4$  and Y are each independently H,  $OR^{10}$ ,  $S(O)_nR^{10}$ ,  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkynyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by  $COOR^8$ , CN,  $C(O)NR^6R^7$ ,  $PO_3R^8$ ,  $SO_3R^8$ , heterocyclic,  $OR^8$ , SH,  $S(O)_nR^9$ ,  $NR^6R^7$ ,  $NH(CO)NR^6R^7$ ,  $NH(CO)OR^9$ ,  $OC(O)OR^9$ , or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted by one or two groups independently selected from  $NR^6R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3R^8$ ,  $OCOR^8$ ,  $PO_3R^8$ , and  $C(O)NR^6R^7$  and heterocyclic;

R<sup>5</sup>, R<sup>6</sup> and R<sup>7</sup> are each independently H, C<sub>1</sub>–C<sub>7</sub> alkyl, C<sub>2</sub>–C<sub>7</sub> alkenyl, C<sub>2</sub>-C<sub>7</sub> alkynyl or C<sub>3</sub>–C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by COOR<sup>8</sup>, CN, OR<sup>8</sup>, NR<sup>8</sup>R<sup>9</sup>, SO<sub>3</sub>R<sup>8</sup>, PO<sub>3</sub>R<sup>8</sup>, halogen, or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted by one or two groups independently selected from COOR<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, and PO<sub>3</sub>R<sup>8</sup> and heterocyclic;

R8 is H, C1-C7 saturated straight chain alkyl or cycloalkyl, CF3 or CH2CF3;

 $R^9$  is  $C_1$ – $C_7$  saturated straight chain alkyl or cycloalkyl;

 $R^{10}$  is  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl, aryl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by  $COOR^8$ , CN,  $C(O)NR^6R^7$ ,  $PO_3R^8$ ,  $SO_3R^8$ , heterocyclic,  $OR^8$ , SH,  $S(O)_nR^9$ ,  $NR^6R^7$ ,  $NH(CO)NR^6R^7$ ,  $NH(CO)OR^9$ , or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from  $NR^6R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3R^8$ ,  $OCOR^8$ ,  $PO_3R^8$ , and  $C(O)NR^6R^7$  and heterocyclic;

Z is  $OR^{11}$ ,  $S(O)_nR^{11}$ ,  $NR^{11}R^{12}$  or  $CHR^{11}R^{12}$ ;

 $R^{11}$  is  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being substituted by  $NR^{13}R^{14}$ ,  $S(O)_nR^{13}$ , or  $OR^{13}$ ;

 $R^{12}$  is hydrogen,  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by  $NR^{13}R^{14}$ ,  $S(O)_nR^{13}$  or  $OR^{13}$ ;

R<sup>13</sup> is SiR<sup>15</sup>R<sup>16</sup>R<sup>17</sup>, C<sub>1</sub>–C<sub>7</sub> alkyl, C<sub>2</sub>–C<sub>7</sub> alkenyl, C<sub>2</sub>–C<sub>7</sub> alkynyl, aryl or C<sub>3</sub>–C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being substituted by one to three groups independently selected from COOR<sup>8</sup>, OR<sup>8</sup>, Si R<sup>15</sup>R<sup>16</sup>R<sup>17</sup>, OR<sup>15</sup>, aryl, and biaryl and heteroaryl, said aryl[[,]] and biaryl and heteroaryl being optionally substituted with one to three groups independently selected from halogen, CF<sub>3</sub>, OR<sup>8</sup>, COOR<sup>8</sup>, NO<sub>2</sub>, and CN;

 $R^{14}$  is H,  $\frac{15}{15}R^{16}R^{17}$ ,  $C_1-C_7$  alkyl,  $C_2-C_7$  alkenyl,  $C_2-C_7$  alkynyl, aryl or  $C_3-C_7$ 

C7 cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by one to three groups independently selected from COOR<sup>8</sup>, OR<sup>8</sup>, Si R<sup>15</sup>R<sup>16</sup>R<sup>17</sup>, OR<sup>15</sup>, aryl, and biaryl and heteroaryl, said aryl[[,]] and biaryl and heteroaryl being optionally substituted with one to three groups independently selected from halogen, CF<sub>3</sub>, OR<sup>8</sup>, COOR<sup>8</sup>, NO<sub>2</sub>, and CN; and or

R<sup>13</sup> and R<sup>14</sup> when taken together with the nitrogen atom to which they are attached may form a 5 – 7 membered heterocyclic ring with one or more heteroatoms selected from O, N and S; said ring being optionally substituted by OR<sup>8</sup>, COOR<sup>8</sup>, or C(O)NR<sup>5</sup>R<sup>6</sup>; and

R15, R16, R17 are each independently is  $C_1$ - $C_7$  alkyl, aryl, benzyl, benzyl, biaryl, heteroaryl, or  $(C_1$ - $C_6)$  alkyl-aryl or  $(C_1$ - $C_6)$  alkyl-heteroaryl, said aryl, benzyl, benzyl, and biaryl being optionally substituted by halogen,  $CF_3$ ,  $OR^8$ ,  $COOR^8$ ,  $NO_2$ , CN, or  $C_1$ - $C_7$  alkyl.

Claim 3. (Currently Amended) A compound of claim 2 wherein  $X_1$  is O or  $S(O)_n$  and Y is  $OR^{10}$  in which  $R^{10}$  is  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl, aryl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by  $COOR^8$ , CN,  $C(O)NR^6R^7$ ,  $PO_3R^8$ ,  $SO_3R^8$ , heterocyclic,  $OR^8$ , SH,  $S(O)_nR^9$ ,  $NR^6R^7$ ,  $NH(CO)NR^6R^7$ ,  $NH(CO)OR^9$ , or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from  $NR^6R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3R^8$ ,  $OCOR^9$ ,  $PO_3R^8$ , and  $C(O)NR^6R^7$  or heterocyclic, said  $R^6$ ,  $R^7$ ,  $R^8$  and  $R^9$  substituents being defined as in claim 2.

Claim 4. (Original) A compound of claim 3 in which R<sup>a</sup> and R<sup>b</sup> taken together represent an oxo (=0) group, or R<sup>a</sup> and R<sup>b</sup> are each independently hydrogen or OH.

Claims 5-6. (Canceled).

Claim 7. (Currently Amended) A compound of claim 3 in which

Z is

$$R^{15}$$
  $R^{18}$  aryl  $-(CH_2)_m$   $O$   $Si$   $R^{16}$   $-Of$   $-(CH_2)_m$   $-N$   $-(CH_2)_p$   $-CH$  aryl  $R^{17}$ 

in which m and p each independently represent an integer of one to six,  $R^{15}$ ,  $R^{16}$ ,  $R^{17}$ -are each independently  $C_1$ - $C_7$ -alkyl or phonyl,  $R^{18}$  is  $C_1$ - $C_7$  alkyl and aryl

represents 
$$X^1$$
 in which  $X^1$  is halogen.

Claim 8. (Canceled).

Claim 9. (Original) A pharmaceutical composition for the inhibition of cytosolic phospholipase A<sub>2</sub> comprising a therapeutically effective amount of a compound of claim 1 and a pharmaceutically acceptable carrier.

Claim 10. (Withdrawn) A method of inhibiting cytosolic phospholipase A<sub>2</sub> in a mammal in need thereof, comprising administering to said mammal a therapeutically effective amount of a compound of claim 1.

Claim 11. (Currently Amended) A compound selected from

$$CI$$

$$CH_3$$

$$CH_3$$

$$CH_3$$

$$CH_3$$

$$CH_3$$

$$CH_3$$

$$CO_2H$$

$$CO_2H$$

$$CO_2H$$

$$CO_2H$$

$$CO_2H$$

$$CO_2H$$

$$CO_2H$$

or a pharmaceutically acceptable salt thereof.

### Claim 12. (Currently Amended) A compound of the formula

$$Z \xrightarrow{X_1 \times X_2 \times X_3 \times X_4} X \xrightarrow{R^a \times R^b \times X_1 \times X_2 \times X_2 \times X_3 \times X_4} X \xrightarrow{I}$$

or a pharmaceutically acceptable salt thereof wherein

 $X_1$  is O, S(O)<sub>n</sub>, co-N-, or -CH<sub>2</sub>-, with the proviso that when  $X_1$  is -CH<sub>2</sub>-,  $R^1$  and  $R^2$  are only halogen;

n is 0, 1 or 2;

R<sup>a</sup> and R<sup>b</sup> when taken together form an oxo (=O) group, or R<sup>a</sup> and R<sup>b</sup> are each independently hydrogen, OH, OCOR<sup>9</sup>, NH<sub>2</sub>, N<sub>3</sub>, NHCOCOR<sup>9</sup>, or F;

X is H;

R<sup>1</sup> and R<sup>2</sup> are each independently H, halogen, OR<sup>9</sup>, or C<sub>1</sub>-C<sub>7</sub> alkyl;

 $R^3$ ,  $R^4$  and Y are each independently H, halogen,  $OR^{10}$ , or  $C_1$ - $C_7$  alkyl, said alkyl being optionally substituted by aryl, said aryl being optionally substituted by one or two  $COOR^8$  groups, with the proviso that not all of  $R^3$ ,  $R^4$  and Y may be the same halogen;

 $R^5$ ,  $R^6$ , and  $R^7$  are each independently hydrogen or  $C_1$ - $C_7$  alkyl, said alkyl being optionally substituted by  $OR^8$ ;

R<sup>8</sup> is H or C<sub>1</sub>-C<sub>7</sub> saturated straight chain alkyl;

R<sup>9</sup> is C<sub>1</sub>-C<sub>7</sub> saturated straight chain alkyl;

 $R^{10}$  is  $C_1$ - $C_7$  alkyl or aryl, said alkyl or aryl group being optionally substituted by  $COOR^8$ ,  $C(O)NR^6R^7$ , heterocyclic, or  $OR^8$ ;

Z is OR<sup>11</sup> or CHR<sup>11</sup>R<sup>12</sup>;

 $R^{11}$  is  $C_1$ - $C_7$  alkyl substituted by  $NR^{13}R^{14}$ ,  $S(O)_nR^{13}$ , or  $OR^{13}$ ;

R<sup>12</sup> is hydrogen;

R<sup>13</sup> is SiR<sup>15</sup>R<sup>16</sup>R<sup>17</sup>-or C<sub>1</sub>-C<sub>7</sub> alkyl, said alkyl substituted by one to three groups independently selected from OR<sup>15</sup> and aryl, said aryl substituted with one halogen;

R14 is C1-C7 alkyl; and

 $R^{15}$ ,  $R^{16}$ , and  $R^{17}$  are each independently is  $C_1$ - $C_7$  alkyl, aryl, or benzhydryl, said aryl and benzhydryl being optionally substituted by halogen.

Claim 13. (Currently Amended) A compound of the formula

$$Z \xrightarrow{R^a R^b} X$$

$$X \xrightarrow{R^1 R^2 R^3 R^4}$$

$$I$$

or a pharmaceutically acceptable salt thereof wherein

 $X_1$  is O,  $S(O)_n$ , or  $-CH_2$ -, with the proviso that when  $X_1$  is  $-CH_2$ -,  $R^1$  and  $R^2$  are only halogen;

n is 0, 1 or 2;

 $R^a$  and  $R^b$  are each independently hydrogen, OH, OCOR $^9$ , NH $_2$ , N $_3$ , NHCOOR $^9$ , NHCOCOR $^9$ , or F;

X is H, CF<sub>3</sub>, OCF<sub>3</sub>, halogen, C<sub>1</sub>–C<sub>7</sub> alkyl, C<sub>2</sub>–C<sub>7</sub> alkenyl, C<sub>2</sub>–C<sub>7</sub> alkynyl or C<sub>3</sub>–C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl or cycloalkyl group being optionally substituted by COOR<sup>8</sup>, CN, C(O)NR<sup>6</sup>R<sup>7</sup>, PO<sub>3</sub>R<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, heterocyclic, OR<sup>8</sup>, SH, S(O)<sub>n</sub>R<sup>9</sup>, NR<sup>6</sup>R<sup>7</sup>, NH(CO)NR<sup>6</sup>R<sup>7</sup>, NH(CO)OR<sup>9</sup>, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from NR<sup>6</sup>R<sup>7</sup>, OR<sup>8</sup>, COOR<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, OCOR<sup>9</sup>, PO<sub>3</sub>R<sup>8</sup>, and C(O)NR<sup>6</sup>R<sup>7</sup> and heterocyclic;

 $R^1$  and  $R^2$  are each independently H, halogen,  $OR^9$ ,  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkynyl,  $C_2$ – $C_7$  alkenyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by  $COOR^8$ , CN,  $C(O)NR^6R^7$ ,  $PO_3R^8$ ,  $SO_3R^8$ , heterocyclic,  $OR^8$ , SH,  $S(O)_nR^9$ ,  $NR^6R^7$ ,  $NH(CO)NR^6R^7$ ,  $NH(CO)OR^9$ ,  $OC(O)OR^9$ , or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted with one or two groups independently selected from  $NR^6R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3R^8$ ,  $OCOR^9$ ,  $PO_3R^8$ , and  $C(O)NR^6R^7$  and heterocyclic;

 $R^3$  and  $R^4$  are each independently H, halogen,  $OR^{10}$ ,  $S(O)_nR^{10}$ ,  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by  $COOR^8$ , CN,  $C(O)NR^6R^7$ ,  $PO_3R^8$ ,  $SO_3R^8$ , heterocyclic,  $OR^8$ , SH,  $S(O)_nR^9$ ,  $NR^6R^7$ ,  $NH(CO)NR^6R^7$ ,  $NH(CO)OR^9$ ,  $OC(O)OR^9$ , or aryl or heteroaryl, said aryl and heteroaryl being optionally substituted by one or two groups independently selected from  $NR^6R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3R^8$ ,  $OCOR^8$ ,  $PO_3R^8$ , and  $C(O)NR^6R^7$  and heterocyclic, with the proviso that not all of  $R^3$ ,  $R^4$  and Y may be the same halogen;

Y is  $OR^{10}$  or  $S(O)_nR^{10}$ ;

R<sup>5</sup>, R<sup>6</sup> and R<sup>7</sup> are each independently H, C<sub>1</sub>–C<sub>7</sub> alkyl, C<sub>2</sub>–C<sub>7</sub> alkenyl, C<sub>2</sub>-C<sub>7</sub> alkynyl or C<sub>3</sub>–C<sub>7</sub> cycloalkyl, said alkyl, alkenyl, alkynyl and cycloalkyl group being optionally substituted by COOR<sup>8</sup>, CN, OR<sup>8</sup>, NR<sup>8</sup>R<sup>9</sup>, SO<sub>3</sub>R<sup>8</sup>, PO<sub>3</sub>R<sup>8</sup>, halogen, or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from COOR<sup>8</sup>, SO<sub>3</sub>R<sup>8</sup>, and PO<sub>3</sub>R<sup>8</sup> and heterocyclic;

R<sup>8</sup> is H, C<sub>1</sub>-C<sub>7</sub> saturated straight chain alkyl or cycloalkyl;

 $R^9$  is  $C_1$ – $C_7$  saturated straight chain alkyl or cycloalkyl;

 $R^{10}$  is  $C_1$ – $C_7$  alkyl,  $C_2$ – $C_7$  alkenyl,  $C_2$ – $C_7$  alkynyl, aryl or  $C_3$ – $C_7$  cycloalkyl, said alkyl, alkenyl, alkynyl, aryl or cycloalkyl group being optionally substituted by  $COOR^8$ , CN,  $C(O)NR^6R^7$ ,  $PO_3R^8$ ,  $SO_3R^8$ , heterocyclie,  $OR^8$ , SH,  $S(O)_nR^9$ ,  $NR^6R^7$ ,  $NH(CO)NR^6R^7$ ,  $NH(CO)OR^9$ , or aryl or heteroaryl, said aryl or heteroaryl being optionally substituted by one or two groups independently selected from  $NR^6R^7$ ,  $OR^8$ ,  $COOR^8$ ,  $SO_3R^8$ ,  $OCOR^8$ ,  $PO_3R^8$ , and  $C(O)NR^6R^7$  or heterocyclie; and

Z is

in which m and p each independently represent an integer of one to six,  $R^{15}$ ,  $R^{16}$ ,  $R^{17}$ -are each independently  $C_1$ - $C_7$  alkyl or phenyl,  $R^{18}$  is  $C_1$ - $C_7$  alkyl and aryl

represents 
$$X^1$$
 in which  $X^1$  is halogen.

### REMARKS/ARGUMENTS

Claims 1-4, 7, and 9-13 are pending in this application.

In the Office Action dated July 8, 2004, the Examiner rejected Claims 1-4, 9, 12, and 13 under 35 U.S.C. §102(b) as being unpatentable over JP 5-222006. Claim 7 was objected to as being dependent upon a rejected base claim.

Reconsideration and allowance of this application are respectfully requested in view of the above amendments and the remarks that follow.

Pursuant to the Examiner's request for an election of a single disclosed species on July 29, 2003, Applicants elected 3-[4-[3-[N-[2-Bis-(4-chlorophenyl)ethyl]-N-methylamino]propyl]phenoxy]-1-(4-carboxyphenoxy)-2-propanone which is Example 2 on page 51 of the specification.

3-[4-[3-[N-[2-Bis-(4-chlorophenyl)ethyl]-N-methylamino]propyl]phenoxy]-1-(4-carboxyphenoxy)-2-propanone

In the July 8, 2004 Office Action, the Examiner states that "Claim 11 will be allowed to the extent it reads on the elected subject matter. Compounds containing Silicon and heterocyclic subject matter should be deleted." Accordingly, Applicants have amended Claims 1, 2, 3, 7,11, 12, and 13 to remove silicon and heterocyclic subject matter which reflects the scope of the generic concept of the elected subject matter. Applicants maintain the right to file divisonal application(s) on non-elected subject matter.

# Rejection of Claims 1-4, 9, 12, and 13 Under 35 U.S.C. §102(b)

The Examiner has rejected Claims 1-4, 9, 12, and 13 under 35 U.S.C. §102(b) as being unpatentable over JP 5-222006. JP'006 teaches compounds containing a

heterocycle. It is Applicants' position that the amendments to Claims 1, 2, 3, 9, 12, and 13, which remove all heterocyclic subject matter, render the rejections moot. Therefore, it is respectfully requested that the rejections to Claims 1-4, 9, 12, and 13 be withdrawn.

### Objection of Claim 7 and Allowance of Claim 11

Claim 7 has been objected to as being dependent upon a rejected base claim, but would be allowable to the extent that it reads on the elected subject matter, if rewritten in independent form including all of limitations of the base claim. The Examiner further states "Note applicants should delete Silicon containing subject matter." Applicants have amended Claim 7 to remove all silicon containing subject matter and respectfully request that the objection be withdrawn.

Claim 11 is allowed to the extent it reads on the elected subject matter. The Examiner further states "Compounds containing Silicon and heterocyclic subject matter should be deleted." Applicants have amended Claim 11 to remove all silicon and heterocyclic subject matter and respectfully request that the claim be allowed.

While Applicants submit that the claims are in condition for allowance and respectfully request the Examiner's reconsideration, a NOTICE OF APPEAL has nevertheless been filed. The Commissioner is hereby authorized to charge any additional fees under 37 CFR §1.17 which may be required, or credit any overpayment, to Account No. 19-3880 in the name of Bristol-Myers Squibb Company.

Respectfully submitted,

Bristol-Myers Squibb Company Patent Department P.O. Box 4000 Princeton, NJ 08543-4000 (203) 677-6997

Date: October 7, 2004

Pamela A. Mingo
Agent for Applicants
Reg. No. 48, 256

	CASE QA211 NP
	CERTIFICATE OF FACSIMILE TRANSMISSION
I hereby certify that this paper (along with any paper redate shown below.  Pamela A. Mingo Type or print name	Tred to as being attached or enclosed) is being facsimile transmitted to the Patent and Trademark Office on the Control of the Control of the Patent and Trademark Office on the Control of the Cont
IN THE UNITED	STATES PATENT AND TRADEMARK OFFICE
IN RE APPLICATION OF Banville et. al APPLICATION NO: 09/848,694	ART UNIT: 1621 EXAMINER: S. Kumar
FILED: 05/03/2001 FOR: Alpha-Amino, -Thio, -Oxo Subst	uted Ketones as Phopholipase Inhibitors
Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450	
	NOTICE OF APPEAL
Sir:  Applicants hereby appeal to the finally rejecting Claims 1-4, 9, 12, and 1	Board of Patent Appeals and Interferences from the Office Action dated July 8, 2004
	No. 19-3880 in the name of Bristol-Myers Squibb Company in the amount of \$330 for

payment of the appeal fee. An additional copy of this paper is here enclosed. The Commissioner is hereby  $\boxtimes$ authorized to charge any additional fees which may be required, or credit any overpayment, to Account No. 19-3880

in the name of Bristol-Myers Squibb Company.

The appeal fee was paid in a previous appeal herein. The examiner re-opened prosecution prior to any decision by the 

Board of Patent Appeals and Interferences. No fee is now due.

Enclosed is a Petition for Extension of Time. 

Respectfully submitted,

Bristol-Myers Squibb Company Patent Department P.O. Box 4000 Princeton, NJ 08543-4000 (203) 677-7268 Date: October 7, 2004

Pamela A. Mingo Agent for Applicants Reg. No. 48,256